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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/582,696	03/09/2007	Stephan Christopher Pleines	70279USPCT	7419

22847 7590 05/25/2010  
SYNGENTA BIOTECHNOLOGY, INC.  
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EXAMINER
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THOMAS, DAVID C

ART UNIT	PAPER NUMBER
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1637

NOTIFICATION DATE	DELIVERY MODE
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05/25/2010

ELECTRONIC

**Please find below and/or attached an Office communication concerning this application or proceeding.**

The time period for reply, if any, is set in the attached communication.

Notice of the Office communication was sent electronically on above-indicated "Notification Date" to the following e-mail address(es):

IP.SBI@syngenta.com

<b>Office Action Summary</b>	<b>Application No.</b> 10/582,696	<b>Applicant(s)</b> PLEINES ET AL.	
	<b>Examiner</b> DAVID C. THOMAS	<b>Art Unit</b> 1637	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

### Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) OR THIRTY (30) DAYS, WHICHEVER IS LONGER, FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

### Status

- 1) ☒ Responsive to communication(s) filed on 01 March 2010.
- 2a) ☐ This action is **FINAL**.                      2b) ☒ This action is non-final.
- 3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

### Disposition of Claims

- 4) ☒ Claim(s) 1-40 is/are pending in the application.
- 4a) Of the above claim(s) 1-11, 20, 21, 30 and 32-39 is/are withdrawn from consideration.
- 5) ☐ Claim(s) \_\_\_\_\_ is/are allowed.
- 6) ☒ Claim(s) 12-19, 22-29, 31 and 40 is/are rejected.
- 7) ☐ Claim(s) \_\_\_\_\_ is/are objected to.
- 8) ☐ Claim(s) \_\_\_\_\_ are subject to restriction and/or election requirement.

### Application Papers

- 9) ☐ The specification is objected to by the Examiner.
- 10) ☐ The drawing(s) filed on \_\_\_\_\_ is/are: a) ☐ accepted or b) ☐ objected to by the Examiner.  
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).  
Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
- 11) ☐ The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

### Priority under 35 U.S.C. § 119

- 12) ☐ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
- a) ☐ All    b) ☐ Some \*    c) ☐ None of:
1. ☐ Certified copies of the priority documents have been received.
2. ☐ Certified copies of the priority documents have been received in Application No. \_\_\_\_\_.
3. ☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

\* See the attached detailed Office action for a list of the certified copies not received.

### Attachment(s)

- |   |   |
|---|---|
| 1) <input checked="" type="checkbox"/> Notice of References Cited (PTO-892)   | 4) <input type="checkbox"/> Interview Summary (PTO-413)<br>Paper No(s)/Mail Date. _____ |
| 2) <input type="checkbox"/> Notice of Draftperson's Patent Drawing Review (PTO-948)   | 5) <input type="checkbox"/> Notice of Informal Patent Application                       |
| 3) <input checked="" type="checkbox"/> Information Disclosure Statement(s) (PTO/SB/08)<br>Paper No(s)/Mail Date <u>12/12/2006</u> . | 6) <input type="checkbox"/> Other: _____  |

## **DETAILED ACTION**

### ***Election/Restrictions***

1. Applicant's election without traverse of Group II, claims 12-19, 22-29, 31 and 40, in the reply filed on March 1, 2010 is acknowledged. Claims 1-11, 20, 21, 30, 32 and 33-39 are withdrawn from further consideration pursuant to 37 CFR 1.142(b) as being drawn to a nonelected invention, there being no allowable generic or linking claim.

### ***Specification***

2. The disclosure is objected to because of the following informalities: The first paragraph on p. 2 of the Specification is incomplete, since it begins in the middle of a sentence and is not a continuation of the last full paragraph on p. 1.

Appropriate correction is required.

### ***Claim Objections***

3. Claims 19 and 22 are objected to because of the following informalities: References to nucleotide sequences in the form of primers are cited in claims 19 and 22. However, these sequences are not properly identified using "SEQ ID NO:" prior to the sequence identifier of the corresponding sequence cited in the Sequence Listing. See MPEP section 2422, particularly with reference to 37 CFR 1.821(d). Appropriate correction is required.

***Claim Rejections - 35 USC § 112***

4. The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

5. Claims 16-18 are rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention. Claims 16-18 recite the limitation "The method of selecting a *Brassica* plant" in claim 12. There is insufficient antecedent basis for this limitation in the claim.

***Claim Rejections - 35 USC § 102***

6. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

7. Claims 12-18, 23-29, 31 and 40 are rejected under 35 U.S.C. 102(b) as being anticipated by Tulsieram et al. (WO 98/56948, cited on IDS).

Tulsieram teaches a method of detecting a *Brassica* plant containing a restorer gene (methods are presented for using molecular markers for determining the genotype of the Ogura restorer gene in *Brassica* plants by the techniques of random amplified

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polymorphic DNA (RAPD) or amplified fragment length polymorphism (AFLP) analysis, see Abstract and p. 6, lines 9-17), comprising the steps of:

a) obtaining a sample from a *Brassica* plant (for analysis of markers by AFLP, plant material was obtained from a winter *B. napus* population, 94CWN2133, p. 28, lines 24-26);

b) detecting in said sample a DNA fragment (AFLP analysis was performed after DNA extraction, using AFLP primers having a single selective nucleotide in a first step amplification, with a second amplification reaction performed using labeled primers, p. 29, lines 7-32; amplification products were resolved by gel electrophoresis, p. 30, lines 1-15; putative markers for the restorer gene were then sequenced, p. 30, lines 17-29), by

ii) at least one marker of bin 2, but none of the markers of bin 3 (AFLP markers included E33/M47 of bin2, but not any bin3 markers, p. 20, lines 18-23 and p. 21, lines 28-30).

With regard to claim 13, Tulsieram teaches a method further comprising selecting said *Brassica* plant, or a part thereof, containing said DNA fragment (the population used for AFLP analysis was derived by crossing homozygous restorer plants with sterile plants from the same population to produce heterozygous offspring, p. 28, lines 26-28).

With regard to claim 14, Tulsieram teaches a method further comprising the step of selfing said *Brassica* plant containing said DNA fragment (the resulting heterozygous offspring were then selfed resulting in a population segregating for the restorer gene, p. 28, lines 28-30).

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With regard to claim 15, Tulsieram teaches a method further comprising the step of crossing said *Brassica* plant with another *Brassica* plant (the winter *B. napus* population, 94CWN2133, was obtained by crossing NW3002 and Bristol canola plants, p. 24, lines 7-9).

With regard to claim 16, Tulsieram teaches a method of selecting a *Brassica* plant, wherein said DNA fragment comprises the BLR1 recombination event (recombination events in fertile offspring plants that result in low glucosinolate *Brassica* plant lines were detected by screening for AFLP markers linked to the restorer gene, with breakpoints indicated by which markers are lost, p. 21, lines 14-26, Table 6 and Figure 3).

With regard to claims 17 and 18, Tulsieram teaches a method of selecting a *Brassica* plant, wherein said marker of bin 2 comprises E33M47, E2M4-1, E3MI-1, E4M14-1, E5M1-2, E5M4-2, or E8MI4-2 or has partial homology to E33M47, E2M4-1, E3MI-1, E4M14-1, E5M1-2, E5M4-2, or E8M14-2 (AFLP markers included E33/M47 of bin2, p. 9, lines 14-19, p. 20, lines 18-23 and p. 21, lines 28-30; note, there is no strict definition in the Specification of "partial homology" and therefore sequences having as few as two common contiguous bases may be considered to share partial homology).

With regard to claim 23, Tulsieram teaches a method for producing a fertile F1 hybrid *Brassica* plant (plants were produced and scored as either male fertile or sterile based on flower phenotype, p. 24, lines 15-17 and p. 28, line 30 to p. 29, line 2) comprising the steps of:

a) determining total glucosinolate content in the male fertile restorer parent comprising the BLR1 recombination event and, optionally, also in the female male sterile CMS parent (plants were determined to be fertile or sterile offspring containing or not containing the fertility restorer gene by testing for glucosinolate levels and also screening for AFLP markers indicating recombination events had occurred, with breakpoints indicating which markers are lost, p. 21, lines 14-26, p. 22, lines 4-8, Table 6 and Figure 3); and

b) crossing the female and male parents to produce F1 hybrid seed (fertile plants, as determined by AFLP analysis, were derived by crossing homozygous restorer plants with sterile plants from the same population to produce heterozygous offspring, p. 28, lines 26-28).

With regard to claim 24, Tulsieram teaches a method for producing a fertile F1 hybrid *Brassica* plant (plants were produced and scored as either male fertile or sterile based on flower phenotype, p. 24, lines 15-17) comprising the steps of:

a) detecting in seed or a plant of the male fertile restorer parent the BLR1 recombination event through marker analysis (recombination events in fertile offspring plants that result in low glucosinolate *Brassica* plant lines were detected by screening for AFLP markers linked to the restorer gene, with breakpoints indicated by which markers are lost, p. 21, lines 14-26, Table 6 and Figure 3); and

b) crossing the female and male parents to produce F1 hybrid seed (fertile plants, as determined by AFLP analysis for Rf markers, were derived by crossing

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homozygous restorer plants with sterile plants from the same population to produce heterozygous offspring, p. 28, lines 26-28).

With regard to claim 25, Tulsieram teaches a method comprising the additional step of detecting in seed or a plant of the restorer parent a DNA fragment through marker analysis (AFLP analysis of markers associated with the fertility restorer gene in *Brassica* plants was performed using AFLP primers having a single selective nucleotide in a first step amplification, with a second amplification reaction performed using labeled primers, p. 29, lines 7-32).

With regard to claim 26, Tulsieram teaches a method comprising the additional step of planting said F1 hybrid seed (heterozygous offspring resulting from crossing homozygous restorer plants and sterile plants were selfed using the resulting hybrid seed and plants, p. 28, lines 26-30).

With regard to claim 27, Tulsieram teaches a method comprising the additional step of harvesting the F2 seed grown from the plant resulting from said F1 seed (seeds from the F2 population 94CWN2133 were harvested and grown in a greenhouse for marker analysis, p. 13, lines 28-29 and p. 28, line 30 to p. 29, line 2).

With regard to claim 28, Tulsieram teaches a method comprising the additional step of determining total glucosinolate content in F2 seed derived from the F1 hybrid plant (plants were determined to be fertile or sterile offspring containing or not containing the fertility restorer gene by testing for glucosinolate levels, p. 21, lines 14-26, p. 22, lines 4-8, Table 6 and Figure 3).



With regard to claim 29, Tulsieram teaches a hybrid F1 Brassica plant produced by the method of claim 26 (heterozygous offspring are produced from crossing homozygous restorer plants and sterile plants, the seeds of which were used for selfing, p. 28, lines 26-30).

With regard to claim 31, Tulsieram teaches a method for producing a Brassica plant containing the BLR1 recombination event (plants were produced and scored as either male fertile, and also recombined in the Rf region to have low glucosinolate levels, or sterile based on flower phenotype, p. 24, lines 15-17 and p. 28, line 30 to p. 29, line 2) comprising the steps of obtaining a Brassica plant containing the BLR1 recombination event (the winter *B. napus* population, 94CWN2133, was used to obtain a homozygous restorer population, p. 24, lines 5-7 and p. 28, lines 24-27), crossing this plant with another Brassica plant to obtain hybrid seed (homozygous restorer plants were crossed with sterile plants from the same population to produce heterozygous offspring, p. 28, lines 26-28), and planting said hybrid seed to produce a Brassica plant containing the BLR1 recombination event (heterozygous offspring resulting from crossing homozygous restorer plants and sterile plants were selfed using the resulting hybrid seed and plants to produce a population segregating for the restorer gene, the seeds of which were used to produce plants that were screened for the restorer gene and scored for the flowering phenotype, p. 28, line 26 to p. 29, line 2).

With regard to claim 40, Tulsieram teaches a method comprising the additional step of detecting in seed or a plant of the restorer parent a DNA fragment through marker analysis (markers associated with the fertility restorer gene in parent *Brassica*

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plants such as NW3002, used to generate the restorer population 94CWN2133, were screened for linkage to the restorer gene, including the marker E33/M47, p. 24, lines 7-9 and Table 6).

***Claim Rejections - 35 USC § 103***

8. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:

(a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negated by the manner in which the invention was made.

9. This application currently names joint inventors. In considering patentability of the claims under 35 U.S.C. 103(a), the examiner presumes that the subject matter of the various claims was commonly owned at the time any inventions covered therein were made absent any evidence to the contrary. Applicant is advised of the obligation under 37 CFR 1.56 to point out the inventor and invention dates of each claim that was not commonly owned at the time a later invention was made in order for the examiner to consider the applicability of 35 U.S.C. 103(c) and potential 35 U.S.C. 102(e), (f) or (g) prior art under 35 U.S.C. 103(a).

10. Claims 19 and 22 are rejected under 35 U.S.C. 103(a) as being unpatentable over Tulsieram et al. (WO 98/56948) in view of Tulsieram et al. (GenBank Accession No. BD192151.1 (2003), herein referred to as Tulsieram (2003)) and further in view of Lowe et al. (Nucleic Acids Res. (1990) 18:1757-1761).

Tulsieram teaches the limitations of claims 12-18, 23-29, 31 and 40, as discussed above.

With regard to claim 19, Tulsieram also teaches a method of detecting a *Brassica* plant (methods are presented for using molecular markers for determining the genotype of the Ogura restorer gene in *Brassica* plants by the techniques of random amplified polymorphic DNA (RAPD) or amplified fragment length polymorphism (AFLP) analysis, see Abstract and p. 6, lines 9-17), further comprising the step of detecting in said sample a DNA fragment obtainable by PCR amplification (AFLP analysis was performed after DNA extraction, using AFLP primers having a single selective nucleotide in a first step amplification, with a second amplification reaction performed using labeled primers, p. 29, lines 7-32).

With regard to claim 22, Tulsieram teaches a method of screening a *Brassica* plant to determine whether it contains the BLR1 recombination event (recombination events that result in low glucosinolate *Brassica* plant lines were screened for AFLP markers linked to the restorer gene, with breakpoints indicated by which markers are lost, p. 21, lines 14-26 and Figure 3), comprising extracting DNA from said *Brassica* plant (DNA extraction was performed using a modified CTAB extraction protocol, p. 29, lines 7-13), subjecting the extraction to a polymerase chain amplification reaction in the presence of DNA fragments represented (AFLP analysis was performed, using AFLP primers having a single selective nucleotide in a first step amplification, with a second amplification reaction performed using labeled primers, p. 29, lines 15-32).

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However, Tulsieram does not teach a method of polymerase chain amplification using primers 1159 (SEQ ID NO:13) and 1160 (SEQ ID NO:14), whereas said DNA fragment is not amplified by the primers PR0004F (SEQ ID NO:19) and PR0004R (SEQ ID NO:20).

Tulsieram (2003) teaches a nucleotide sequence used for genotype determination of the Ogura Rf gene in *Brassica Napus* by molecular marker analysis, and includes a 143-base sequence that contains primer binding sites for SEQ ID NO: 13 (positions 2-24) and SEQ ID NO: 14 (positions 138-116) of the instant invention, but not for SEQ ID NOS: 19 and 20 (see search results below):

```

RESULT 2
BD192151
LOCUS      BD192151                143 bp    DNA        linear    PAT 17-JUL-
2003
DEFINITION Use of molecular markers for genotype determination of the ogura
Rf
            gene in brassica napus.
ACCESSION  BD192151
VERSION    BD192151.1  GI:33001890
KEYWORDS   JP 2002512523-A/14.
SOURCE     Aspergillus oryzae
ORGANISM   Aspergillus oryzae
            Eukaryota; Fungi; Dikarya; Ascomycota; Pezizomycotina;
            Eurotiomycetes; Eurotiomycetidae; Eurotiales; Trichocomaceae;
            mitosporic Trichocomaceae; Aspergillus.
REFERENCE  1 (bases 1 to 143)
AUTHORS    Tulsieram,L.K. and Riddell,C.M.
TITLE      Use of molecular markers for genotype determination of the ogura
Rf
            gene in brassica napus
JOURNAL    Patent: JP 2002512523-A 14 23-APR-2002;
            PIONEER HI BRED INTERNATIONAL INC
COMMENT    PN    JP 2002512523-A/14
            PD    23-APR-2002
            PF    10-JUN-1998 JP 1999501170
            PR    10-JUN-1997 CA          2206673
            PI    LOMAS K TULSIERAM,CHRISTINE M RIDDELL
            PC    C12Q1/68,A01H1/04
            CC    Strandedness: Double;
            CC    Topology: Linear;
            CC    /note= 'E33M47 (or E33XM47AI) '
            FH    Key          Location/Qualifiers.

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FEATURES                      Location/Qualifiers  
     source                      1. .143  
                                 /organism="Aspergillus oryzae"  
                                 /mol\_type="genomic DNA"  
                                 /db\_xref="taxon:5062"

## ORIGIN

Query Match                      100.0%;    Score 23;    DB 9;    Length 143;  
 Best Local Similarity    100.0%;  
 Matches    23;    Conservative    0;    Mismatches    0;    Indels    0;    Gaps  
 0;

Qy                      1 TAACAAAATAGAGGGAGAGGATG 23  
                                 |||||  
 Db                      2 TAACAAAATAGAGGGAGAGGATG 24

## RESULT 2

BD192151/c

LOCUS                      BD192151                      143 bp                      DNA                      linear                      PAT 17-JUL-  
 2003

DEFINITION    Use of molecular markers for genotype determination of the ogura  
 Rf

gene in brassica napus.

ACCESSION    BD192151

VERSION      BD192151.1    GI:33001890

KEYWORDS     JP 2002512523-A/14.

SOURCE       Aspergillus oryzae

ORGANISM     Aspergillus oryzae

Eukaryota; Fungi; Dikarya; Ascomycota; Pezizomycotina;  
 Eurotiomycetes; Eurotiomycetidae; Eurotiales; Trichocomaceae;  
 mitosporic Trichocomaceae; Aspergillus.

REFERENCE    1 (bases 1 to 143)

AUTHORS      Tulsieram,L.K. and Riddell,C.M.

TITLE                      Use of molecular markers for genotype determination of the ogura  
 Rf

gene in brassica napus

JOURNAL       Patent: JP 2002512523-A 14 23-APR-2002;  
 PIONEER HI BRED INTERNATIONAL INC

COMMENT      PN    JP 2002512523-A/14

PD    23-APR-2002

PF    10-JUN-1998 JP 1999501170

PR    10-JUN-1997 CA                      2206673

PI    LOMAS K TULSIERAM,CHRISTINE M RIDDELL

PC    C12Q1/68,A01H1/04

CC    Strandedness: Double;

CC    Topology: Linear;

CC    /note= 'E33M47 (or E33XM47AI)'

FH    Key                      Location/Qualifiers.

FEATURES                      Location/Qualifiers

source                      1. .143

/organism="Aspergillus oryzae"

/mol\_type="genomic DNA"

```
                                /db_xref="taxon:5062"
ORIGIN

Query Match          100.0%;  Score 23;  DB 9;  Length 143;
Best Local Similarity 100.0%;
Matches    23;  Conservative    0;  Mismatches    0;  Indels    0;  Gaps
0;

Qy          1 CAAGATTATAGCTACCTAACAGG 23
            |||
Db          138 CAAGATTATAGCTACCTAACAGG 116
```

It would have been prima facie obvious to one having ordinary skill in the art at the time the invention was made to design amplification primers for detection of markers for genotyping the Ogura Rf gene in *Brassica napus* based on the sequence taught by Tulsieram (2003) since this sequence is known to contain markers associated with the fertility restorer gene. Thus, an ordinary practitioner would have been motivated to use such a sequence in order to design primers that are specific for a marker that is associated with the presence of the restorer gene in *Brassica* plants. The ability to detect such markers allows breeders to detect the presence of the restorer gene in offspring plants at an early stage of development instead of having to wait until the plants mature to determine whether the plants produce normal flowers and pollen, an indication of fertile plants (Tulsieram, p. 2, lines 19-23).

In the recent court decision *KSR International Co. v. Teleflex Inc.*, 82 127 SCt 1727 (2007), the U.S. Supreme Court determined that if the combination of the claimed elements was “obvious to try” by a person of ordinary skill, this might show that such a combination was obvious under §103. Regarding “obvious to try”, the Court stated: “A person of ordinary skill is also a person of ordinary creativity, not an automaton. The same constricted analysis led the Court of Appeals to conclude, in error, that a patent claim cannot be proved obvious merely by showing that the combination of

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elements was "obvious to try." *Id.*, at 289 (internal quotation marks omitted). When there is a design need or market pressure to solve a problem and there are a finite number of identified, predictable solutions, a person of ordinary skill has good reason to pursue the known options within his or her technical grasp. If this leads to the anticipated success, it is likely the product not of innovation but of ordinary skill and common sense. In that instance the fact that a combination was obvious to try might show that it was obvious under §103."

Since the claimed primers simply represent structural homologs of the oligonucleotides taught by Tulsieram (2003), which are 100% derived from sequences expressly suggested by the prior art of Tulsieram as useful for primers for the detection and genotyping of *Brassica* plants containing the fertility restorer gene and concerning which a biochemist of ordinary skill would attempt to obtain alternate compounds with improved properties, the claimed primers are *prima facie* obvious over the cited references in the absence of secondary considerations.

With regard to the issue of equivalence of the primers, MPEP 2144.06 notes "Substituting equivalents known for the same purpose. In order to rely on equivalence as a rationale supporting an obviousness rejection, the equivalency must be recognized in the prior art, and cannot be based on applicant's disclosure or the mere fact that the components at issue are functional or mechanical equivalents. An express suggestion to substitute one equivalent component or process for another is not necessary to render such substitution obvious. In re Fout , 675 F.2d 297, 213 USPQ 532 (CCPA 1982)."

With regard to the issue of reasonable expectation of success in using such equivalents at the time the invention was made, as shown in *Lowe*, one of skill in the art was clearly aware of the factors involved in designing amplification primers from a

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known sequence, and would have routinely and predictably designed any such primers. Specifically, Lowe teaches a computer program based on a set of rules which take into account both the sequence of the primers and the amplified region of DNA, such that primer-to-target hybridization is enhanced, while facilitating attainment of full-length extension products by minimizing non-specific product formation and self-priming (see Abstract and p. 1757, column 2, line 33 to p. 1758, column 1, line 41). The program has been tested on a variety of gene products for RT-PCR, for both total and cytoplasmic RNA samples prepared by several different methods (Lowe, p. 1758, column 2, last 2 lines). "Experimental testing has shown that all the amplification products specified by these primers are of the predicted size and also hybridize with the appropriate cDNA or internal oligonucleotide probe" (Lowe, p. 1769, column 2, line 4-8).

### ***Conclusion***

11. Claims 12-19, 22-29, 31 and 40 are rejected. No claims are allowable.

### ***Correspondence***

12. Any inquiry concerning this communication or earlier communications from the examiner should be directed to David C. Thomas whose telephone number is 571-272-3320 and whose fax number is 571-273-3320. The examiner can normally be reached on 5 days, 9-5:30.



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If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Gary Benzion can be reached on 571-272-0782. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

/David C Thomas/  
Examiner, Art Unit 1637

/Kenneth R Horlick/  
Primary Examiner, Art Unit 1637